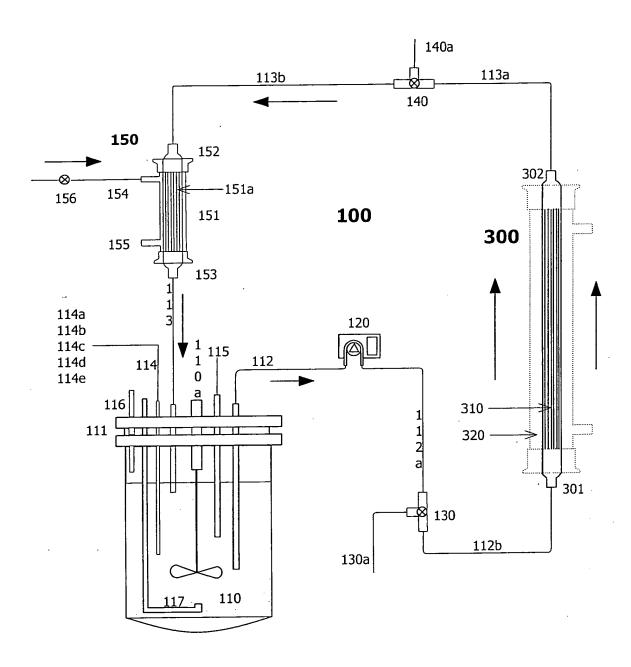
Apparatus and Method for Producing and Using High-Density Cells and Product Therefrom



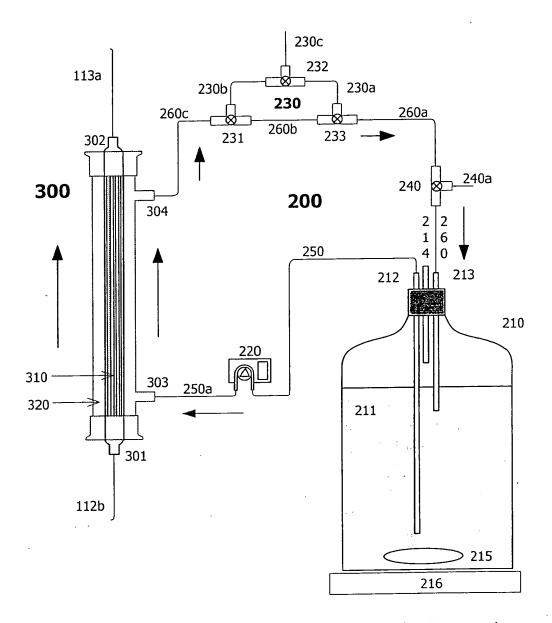


Figure 2



Bioreactor

Figure 3



Media Reservoir

Figure 4

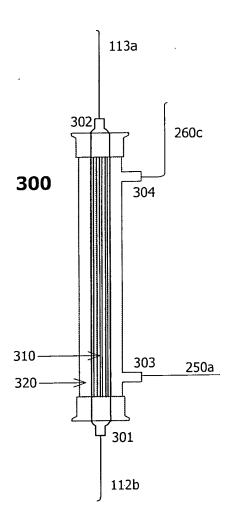


Figure 2. Growth of Insect Cells in a High-Density Dialysis Bioreactor with In-Line Oxygen Sparging.

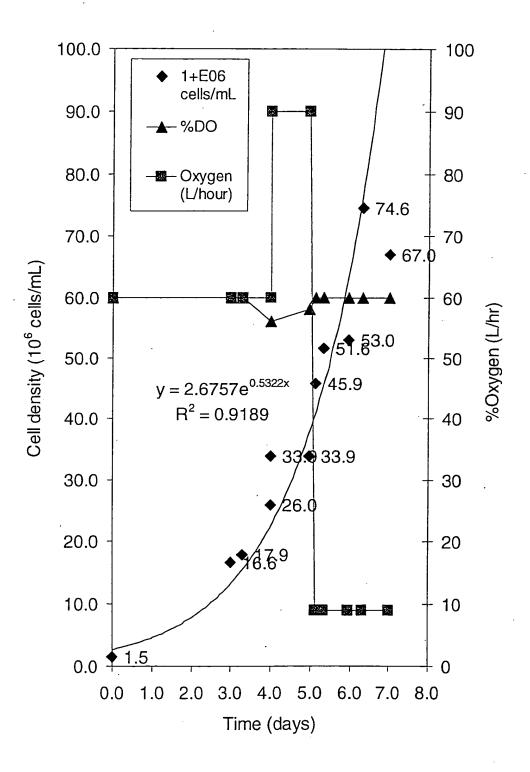
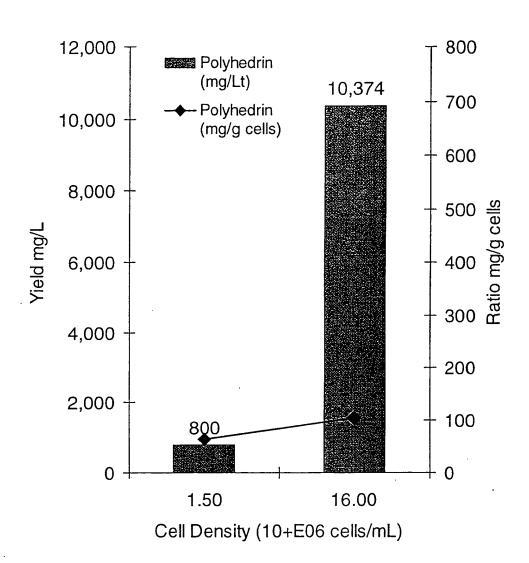
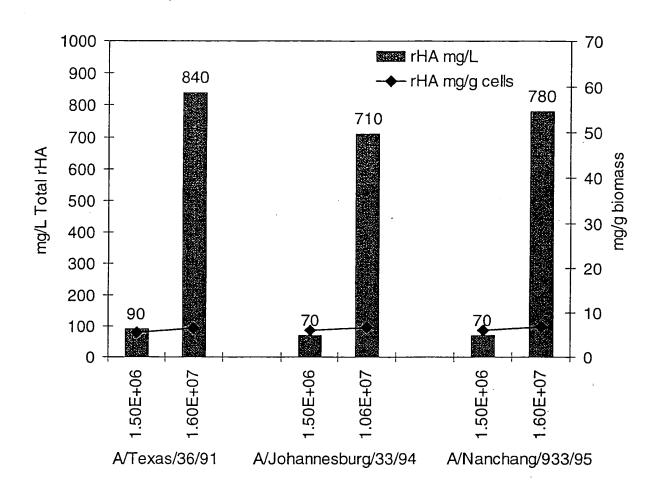
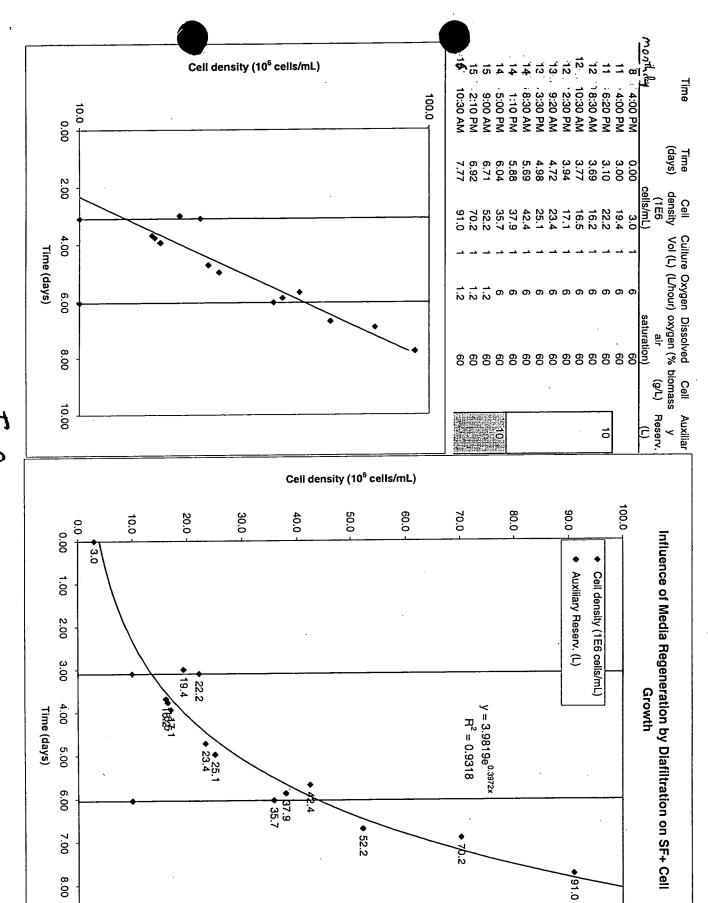


Figure 1. Yields of AcNPV Polyhedrin Protein in Standard and High-Density Cultures.



Yields of Recombinant Hemagglutinin from three Strains of Viral Influenza in Standard and High-Density Cultures.





9.00



HD Bioreactor Diagram Legend

100 Cell Culturing Loop

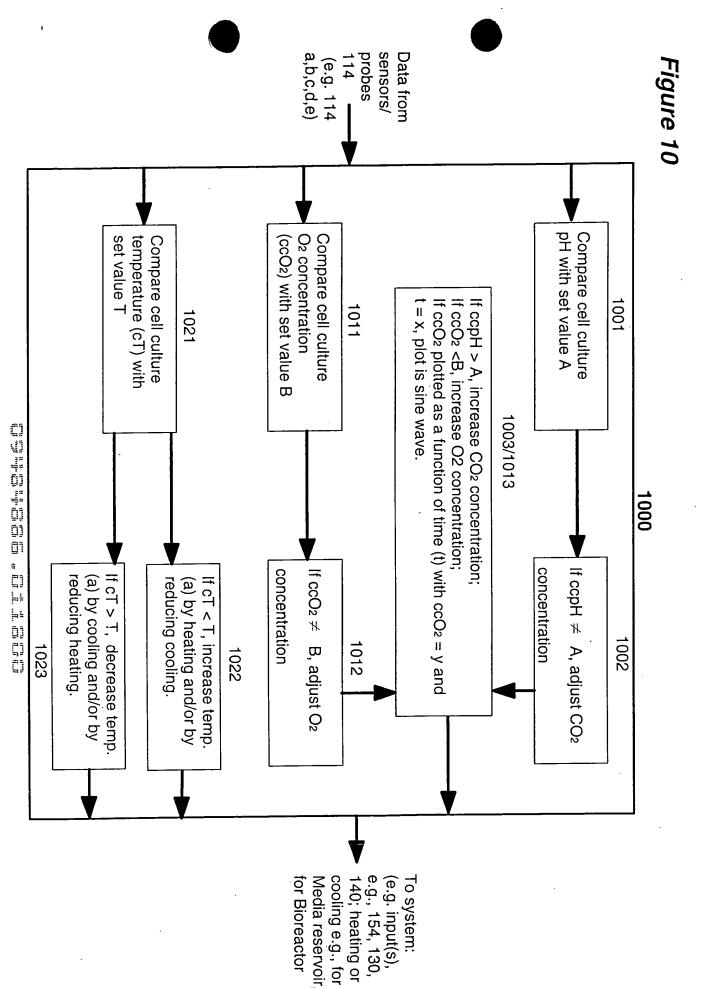
310

320

	110		Stirred-tank Bioreactor
	110	111	Bioreactor headplate
		112	Cell take-up & cell take-up lines (112a – b)
		113	Cell return & cell return lines (113a – b)
		114	Probe ports: multiple (114a – 114e)
			Sampling port
		115	Vent tube
		116	
	400	117	Sparging tube
	120		Pump Three way yelve 8 three way yelve line (120a)
	130		Three-way valve & three-way valve line (130a)
	140		Three-way valve & three-way valve line (140a)
	150		Oxygenation Loop
1.7 mg ng 1.200	•	151	Oxygenator & oxygenator Lumen (151a)
		152	Lumen input
		153	Lumen outflow
		154	Gas input
		155	Gas ouput
		156	Selenoid
		·•	
200	Medium	Replenishment Loop	
- - - -		•	
Ō)	210		Media reservoir
0]		211	Media container
1 1		212	Media take-up
		213	Media return
Cj		214	Vent tube
þ.		215	Magnetic stir bar
<u>L</u> a		216	Variable speed magnetic motor
1 1 1 1 1 1 1 1	220	2.0	Pump
	230	·	"Extraction" loop and "extraction" loop lines (230a - c)
Ēj	200	231	Three-way valve: pass-through or bypass in-line analysis
		232	Three-way valve: collection or sampling
		233	Three-way valve: pass through or return
	240	233	Three-way valve – sampling & three-way valve – sampling line (240a)
	_ · ·		Media take-up lines (250 & 250a)
	250		Media return lines (260 & 260a – c)
	260		Miedia letuiti iilles (200 & 200
300	Hollow	Fiber Dialysis Device	
500	1 IOHOW	i iboi biaijoio botioo	
		301	Lumen input
		302	Lumen outflow
		303	Extra-lumenal input
		304	Extra-lumenal outflow
	040	OW 1	Luman angga

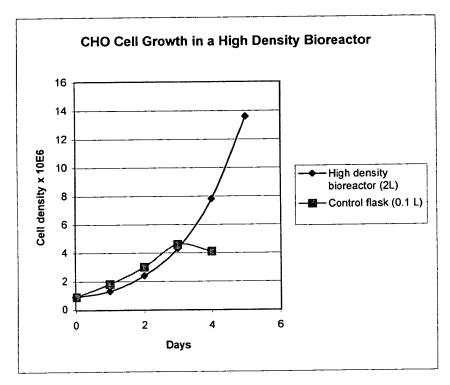
Lumen space

Extra-lumen space



Cell/ml x 10E6

Days	High density bioreactor (2L)	Control flask (0.1 L)
0	0.9	0.9
1	1.3	1.8
2	2.4	3.0
3	4.3	4.6
4	7.8	4.1
5	13.6	



F16.11